What is Claimed:

Par

1. A compound of the formula

Q is selected from the group consisting of

P/8

Re

P/8

R₉COO R₈

R₁₀O R₈

R₉O

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G is selected from the group consisting of alkyl, substituted alkyl, substituted or unsubstituted axyl, heterocyclo,

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W is O or NR₁₅;

X is O or H, H;

Y is selected from the group consisting of O; H, OR $_{16}$; OR $_{17}$, OR $_{17}$; NOR $_{18}$; H, NOR $_{19}$; H, NR $_{20}$ R $_{21}$; H, H; or CHR $_{22}$;OR $_{17}$ OR $_{17}$ can be a cyclic ketal;

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 Z_1 , and Z_2 are selected from the group consisting of CH_2 , O, NR_{23} , S, or SO_2 , wherein only one of Z_1 and Z_2 can be a heteroatom;

 B_1 and B_2 are selected from the group consisting of OR_{24} , or $OCOR_{25}$, or $O_2CNR_{26}R_{27}$; when B_1 is H and Y is OH, H they can form a six-membered ring ketal or acetal;

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D is selected from the group consisting of $NR_{28}R_{29}$, $NR_{30}COR_{31}$ or saturated heterocycle;

R₁, R₂, R₃, R₄, R₅, R₆, R₇, R₁₃, R₁₄, R₁₈, R₁₉, R₂₀, R₂₁, R₂₂, R₂₆, and R₂₇ are selected from the group H, alkyl, substituted alkyl, or aryl and when R₁ and R₂ are alkyl can be joined to form a cycloalkyl; R₃ and R₄ are alkyl can be joined to form a cycloalkyl;

R9, R₁₀, R₁₆, R₁₇, R₂₄, R₂₅, and R₃₁ are selected from the group H, alkyl, or substituted alkyl;

R₈, R₁₁, R₁₂, R₂₈, R₃₀, R₃₂, R₃₃, and R₃₀ are selected from the group consisting of H, alkyl, substituted alkyl, aryl, substitued aryl, cycloalkyl, or heterocyclo;

R₁₅, R₂₃ and R₂₉ are selected from the group consisting of H, alkyl, substituted alkyl, aryl, substituted aryl, cycloalkyl, heterocyclo, R₃₂C=O, R₃₃SO₂, hydroxy, O-alkyl or O-substituted alkyl, the pharmaceutically acceptable salts thereof and any hydrates, solvates or geometric, optical and stereoisomers thereof, with the proviso that compounds wherein

W and X are both O; and

 R_1 , R_2 , R_7 , are H; and

R₃, R₄, R₆, are methyl; and

Rg, is H or methyl; and

 Z_1 , and Z_2 , are CH_2 ; and

G is 1-methyl-2-(substituted-4-thiazolyl)ethenyl; and

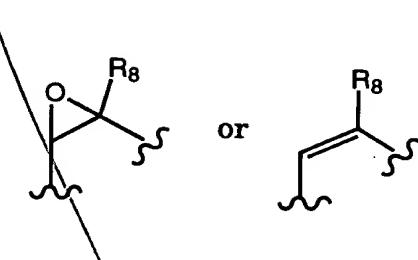
Q is as defined above

25 are excluded.

1,0630

2. The compound of claim 1 wherein

Q is



Cont 5

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X is O;

Y is O;

Z₁, and Z₂, are CH₂; and

W is NR₁₅.

3. A compound selected from the group consisting of:

[1S-[1R*,3R*(E),7R*,10S*,11R*,12R*,16S*]]-7,11-Dihydroxy-8,8,10,12,16pentamethyl-3-[1-methyl-2-(2-methyl- 4-thiazolyl)ethenyl]-4,13,17trioxabicyclo[14.1.0]heptadecane-5,9-dione;

[1S-[1R*,3R*(E),7R*,10S*,11R*,12R*,16S*]]-7,11-Dihydroxy-8,8,10,12tetramethyl-3-[1-methyl-2-(2-methyl-4-thiazolyl)ethenyl]-4,13,17trioxabicyclo[14.1.0]heptadecane-5,9-dione;

[4S-[4R*,7S*,8R*,9R*,15R*(E)]]-4,8-Dihydroxy-5,5,7,9,13-pentamethyl-16-[1-methyl-2-(2-methyl- 4-thiazolyl)ethenyl]-1,10-dioxa-13-

cyclohexadecene-2,6-dione;

[4S-[4R*,7S*,8R*,9R*,15R*(E)]]-4,8-Dihydroxy-5,5,7,9-tetramethyl-16-[1-methyl-2-(2-methyl- 4-thiazolyl)ethenyl]-1,10-dioxa-13-cyclohexadecene-2,6-dione;

[1S-[1R*,3R*(E),7R*,10S*,11R*,12R*,16S*]]-7,11-Dihydroxy-8,8,10,12,16-pentamethyl-3-[1-methyl-2-(2-methyl- 4-thiazolyl)ethenyl]-4,14,17-trioxabicyclo[14.1.0]heptadecane-5,9-dione; λ

30 [1S-[1R*,3R*(E),7R*,10S*,11R*,12R*,16S*]]-7,11-Dihydroxy-8,8,10,12-tetramethyl-3-[1-methyl-2-(2-methyl-4-thiazolyl)ethenyl,-4,14,17-trioxabicyclo[14.1.0]heptadecane-5,9-dione;



The first

[4S-[4R*,7S*,8R*,9R*,15R*(E)]]-4,8-Dihydroxy-5,5,7,9,13-pentamethyl-16-pentamet11-methyl-2-(2-methyl-4-thiazolyl)ethenyl]-1,11-dioxa-13cyclonexadecene-2,6-dione;

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[4S-[4R*,7S*,8R*,9R*,15R*(E)]]-4,8-Dihydroxy-5,5,7,9-tetramethyl-16-[1-16]methyl-2-(2-methyl-4-thiazolyl)ethenyl]-1,11-dioxa-13-cyclohexadecene-2,6-dione;

[1S-[1R*,3R*(E),7R*,10S*,11R*,12R*,16S*]]-7,11-Dihydroxy-8,8,10,12,16-10 pentamethyl-3-[1-methyl-2-(2-methyl-4-thiazolyl)ethenyl]-4,17dioxabicyclo[14.1.0]heptadecane-9-one;

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[1S-[1R*,3R*(E),7R*,10S*,11R*,12R*,16S*]]-7,11-Dihydroxy-8,8,10,12tetramethyl-3-[1-methyl-2-(z-methyl-4-thiazolyl)ethenyl]-4,17dioxabicyclo[14.1.0]heptadecane-9-one;

[1S-[1R*,3R*(E),7R*,10S*,11R*,12R*,16S*]]-7,11-Dihydroxy-3,8,8,10,12,16hexamethyl-3-[1-methyl-2-(2-methyl-4-thiazolyl)ethenyl]-4,17-

dioxabicyclo[14.1.0]heptadecane-5,9-dione; 20

> [1S-[1R*,3R*(E),7R*,10S*,11R*,12R*,16S*]]-7,11-Dihydroxy-3,8,8,10,12pentamethyl-3-[1-methyl-2-(2-methyl-4-thiazolyl)ethenyl]-4,17dioxabicyclo[14.1.0]heptadecane-5,9-dione;

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[4S-[4R*,7S*,8R*,9R*,15R*(E)]]-4,8-Dihydroxy-5,5,7,9,13,16-hexamethyl-16-[1-methyl-2-(2-methyl- 4-thiazolyl)ethenyl]-1-oxa-13-cyclohexadecene-2,6-dione;

[4S-[4R*,7S*,8R*,9R*,15R*(E)]]-4,8-Dihydroxy-5,5,7,9,16-pentamethyl-16-30 [1-methyl-2-(2-methyl-4-thiazolyl)ethenyl]-1-oxa-13-cyclohexaqecene-2,6dione;

[1S-[1R*,3R*(E),7R*,10S*,11R*,12R*,16S*]]-7,11-Dihydroxy-8,8,10,12,16pentamethyl-3-[1-methyl-2-(2-methyl-4-thiazolyl)ethenyl]-4,17-35 dioxabicyclo[14.1.0]heptadecane-5,9-dione;

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[1S-[1R*,3R*(E),7R*,10S*,11R*,12R*,16S*]]-7,11-Dihydroxy-6,8,8,10,12-pentamethyl-3-[1-methyl-2-(2-methyl-4-thiazolyl)ethenyl]-4,17-dioxabiçyclo[14.1.0]heptadecane-5,9-dione;

 $[1S-[1R^*,3R^*(E),7R^*,10S^*,11R^*,12R^*,16S^*]]-7,11-Dihydroxy-8,8,10,12,16-pentamethyl-3-[1-methyl-2-(2-methyl-4-thiazolyl)ethenyl]-4-aza-17-oxabicyclo[14.1.0]heptadecane-5,9-dione;$

10 [1S-[1R*,3R*(E),7R*,\0S*,11R*,12R*,16S*]]-7,11-Dihydroxy-8,8,10,12-tetramethyl-3-[1-methyl-2-(2-methyl- 4-thiazolyl)ethenyl]-4-aza-17-oxabicyclo[14.1.0]heptauccane-5,9-dione;

[4S-[4R*,7S*,8R*,9R*,15R*(E)]]-4,8-Dihydroxy-5,5,7, 9,13-pentamethyl-16-[1-methyl-2-(2-methyl- 4-thiazolyl)ethenyl]-1-aza-13-cyclohexadecene-2,6-dione;

[4S-[4R*,7S*,8R*,9R*,15R*(E)]]-4,8-Dihydroxy-5,5,7, 9-tetramethyl-16-[1-methyl-2-(2-methyl- 4-thiazolyl)ethenyl] 1-aza-13-cyclohexadecene-2,6-dione;

[1S-[1R*,3R*(E),7R*,10S*,11R*,12R*,16S*]]-7,1\[-Dihydroxy-4,8,8,10,12,16-hexamethyl-3-[1-methyl-2-(2-methyl- 4-thiazolyi) ethenyl]-4-aza-17-oxabicyclo[14.1.0]heptadecane-5,9-dione;\[-]

[1S-[1R*,3R*(E),7R*,10S*,11R*,12R*,16S*]]-7,11-Dihydkoxy-4,8,8,10,12-pentamethyl-3-[1-methyl-2-(2-methyl- 4-thiazolyl)ethenyl-4-aza-17-oxabicyclo[14.1.0]heptadecane-5,9-dione;

[4S-[4R*,7S*,8R*,9R*,15R*(E)]]-4,8-Dihydroxy-1,5,5,7, 9,13-hexamethyl-16-[1-methyl-2-(2-methyl-4-thiazolyl)ethenyl]-1-aza-13-cyclohexadecene-2,6-dione;

[4S-[4R*,7S*,8R*,9R*,15R*(E)]]-4,8-Dihydroxy-1,5,5,7, 9-pentamethyl-1&- [1-methyl-2-(2-methyl- 4-thiazolyl)ethenyl]-1-aza-13-cyclohexadecene-2,6 dione;

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[1S-[1R*,3R*(E),7R*,10S*,11R*,12R*,16S*]]-7,11-Dihydroxy-8,8,10,12,16-pentamethyl-3-[1-methyl-2-(2-methyl- 4-thiazolyl)ethenyl]-13-aza-4,17-dioxabicyclo[14.1.0]heptadecane-5,9-dione;

[1S-[1R*\3R*(E),7R*,10S*,11R*,12R*,16S*]]-7,11-Dihydroxy-8,8,10,12-tetrameth χ [-3-[1-methyl-2-(2-methyl-4-thiazolyl)ethenyl]-13-aza-4,17-dioxabicyclo χ [4.1.0]heptadecane-5,9-dione;

10 [4S-[4R*,7S*,8R*,9R*,15R*(E)]]-4,8-Dihydroxy-5,5,7,9,13-pentamethyl-16-[1-methyl-2-(2-methyl- 4-thiazolyl)ethenyl]-10-aza-1-oxa-13cyclohexadecene-2,6-dione;

[4S-[4R*,7S*,8R*,9R*,15R*(E)]]-4,8-Dihydroxy-5,5,7,9-tetramethyl-16-[1-methyl-2-(2-methyl-4-thiazolyl)ethenyl]-10-aza-1-oxa-13-cyclohexadecene-2,6-dione;

[1S-[1R*,3R*(E),7R*,10S*,11R*,12R*,16S*]]-7,11-Dihydroxy-8,8,10,12,16-pentamethyl-3-[1-methyl-2-(2-methyl-4-thiazolyl)ethenyl]-14-aza-4,17-dioxabicyclo[14.1.0]heptadecane-5,9-dione;

[1S-[1R*,3R*(E),7R*,10S*,11R*,12R*,16S*]]-7,11-Dihydroxy-8,8,10,12-tetramethyl-3-[1-methyl-2-(2-methyl-4-thiazoiyl)ethenyl]-14-aza-4,17-dioxabicyclo[14.1.0]heptadecane-5,9-dione;

[4S-[4R*,7S*,8R*,9R*,15R*(E)]]-4,8-Dihydroxy-5,5,7,9,13-pentamethyl-16-[1-methyl-2-(2-methyl- 4-thiazolyl)ethenyl]-11-aza i oxa-13cyclohexadecene-2,6-dione;

[4S-[4R*,7S*,8R*,9R*,15R*(E)]]-4,8-Dihydroxy-5,5,7,9-tetramethyl-16-[1-methyl-2-(2-methyl- 4-thiazolyl)ethenyl]-11-aza-1-oxa-13-cyclohexadecene-2,6-dione;



[1S-[1R*,3R*,7R*,10S*,11R*,12R*,16S*]]-N-Phenyl-7,11-dihydroxy-8,8,10,12-tetramethyl-5,9-dioxo-4,17-dioxabicyclo[14.1.0]heptadecane-3-carboxamide;

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[4S-[4R*,7S*,8R*,9R*,15R*]]-N-Phenyl-4,8-dihydroxy-5,5,7,9,13-pentamethyl-2,6-dioxo-1-oxa-13-cyclohexadecene-16-carboxamide;

[4S-[4R*,7S*,8R*,9R*,15R*]]-N-Phenyl-4,8-dihydroxy-5,5,7,9-tetramethyl-2,6-dioxo-1-oxa-13-cyclohexadecene-16-carboxamide;

[1S-[1R*,3R*(E),7R*,10S*,11R*,12R*,16S*]]-7,11-Dihydroxy-8,8,10,12,16-pentamethyl-3-[1-methyl-2-(2-methyl-4-thiazolyl)cyclopropyl]-4,17-dioxabicyclo[14.1.0]heptadecane-5,9-dione//

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[1S-[1R*,3R*(E),7R*,10S*,11R*,12R*,16S*]]-7,11-Dihydroxy-8,8,10,12-tetramethyl-3-[1-methyl-2-(2-methyl- 4-thiazolyl)cyclopropyl]-4,17-dioxabicyclo[14.1.0]heptadecane-5,9-dione/;

[4S-[4R*,7S*,8R*,9R*,15R*(E)]]-4,8-Dihydroxy-5,5,7,9,13-pentamethyl-16-[1-methyl-2-(2-hydroxymethyl-4-thiazolyl)ethenyl]-1-aza-13(Z)cyclohexadecene-2,6-dione;

and the pharmaceutically acceptable salts, solvates and hydrates thereof.

ph3

- 4. A method of treating cancer in a patient which comprises providing an effective amount of a compound of claim 1 to said patient.
- 5. A method of treating hyperproliferative cellular disease in a patient which comprises providing an effective amount of a compound of claim 1 to said patient.



6. A method of providing an antiangiogenic effect in a patient in need of said treatment which comprises providing an effective amount of a compound of claim 1 to said patient.

add